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INTERNATIONAL PRELIMINARY REPORT ON PATENTABILITY

(Chapter I of the Patent Cooperation Treaty)

(PCT Rule 44bis)

Applicant's or agent's file reference C1-A0505P	FOR FURTHER ACTION	See item 4 below		
International application No. PCT/JP2006/309890	International filing date (day/month/year) 18 May 2006 (18.05.2006)	Priority date (day/month/year) 18 May 2005 (18.05.2005)		
International Patent Classification (8th edition unless older edition indicated) See relevant information in Form PCT/ISA/237				
Applicant THE UNIVERSITY OF TOKUSHIMA				

1.	This international preliminary report on patentability (Chapter I) is issued by the International Bureau on behalf of the International Searching Authority under Rule 44 <i>bis</i> .1(a).			
2.	This REPORT consists of a total of 5 sheets, including this cover sheet.			
	In the attached sheets, any reference to the written opinion of the International Searching Authority should be read as a reference to the international preliminary report on patentability (Chapter I) instead.			
3.	This report contains indications relating to the following items:			
	Box No. I	Basis of the report		
	Box No. II	Priority		
	Box No. III	Non-establishment of opinion with regard to novelty, inventive step and industrial applicability		
	Box No. IV	Lack of unity of invention		
	Box No. V	Reasoned statement under Article 35(2) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement		
	Box No. VI	Certain documents cited		
	Box No. VII	Certain defects in the international application		
	Box No. VIII	Certain observations on the international application		
4.		ommunicate this report to designated Offices in accordance with Rules 44bis.3(c) and 93bis.1 but makes an express request under Article 23(2), before the expiration of 30 months from the priority		
		Date of issuance of this report 19 November 2007 (19.11.2007)		

Authorized officer

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Facsimile No. +41 22 338 82 70 Form PCT/IB/373 (January 2004)

The International Bureau of WIPO 34, chemin des Colombettes

1211 Geneva 20, Switzerland

PATENT COOPERATION TREATY

From the		I AL SEARCI IIN	G AUTHORI	TY			MANS	
То:							PCT	ATTON
					WRITTEN OPINION OF THE INTERNATIONAL SEARCHING AUTHORITY			
					(PCT Rule 43bis.1)			
					Date of mailing (day/month/ye			
Applica	nt's or a	gent's file referenc	e		FOR FURT	HER ACTIO	ON	
C1-	A050)5P				See p	aragraph 2 below	
		plication No. 2006/3098		International filing date (
				national classification and		,		
A61	P1/1	-		A61K38/21(2 1P3/10(2006			21/04(2006.01) 6(2006.01),	۱,
Applica								
THE	UNI	VERSITY	OF TOK	USHIMA				
<u></u>								
I.	This o	pinion contains in	dications relati	ng to the following items	:			
	Box No. I Basis of the opinion							
		Box No. II	Priority					
		Box No. III	Non-establish	nment of opinion with reg	gard to novelty, i	nventive ste	p and industrial applicability	,
		Box No. IV	Lack of unity	ofinvention				
		Box No. V			ale 43bis.1(a)(i) with regard to novelty, inventive step or industrial splanations supporting such statement			
		Box No. VI	Certain docu	ments cited				
	\sqcup	Box No. VII	Certain defec	ts in the international app	lication			
	Ш	Box No. VIII	Certain obser	vations on the internation	nal application			
2.	FURT	HER ACTION						
	If a demand for international preliminary examination is made, this opinion will be considered to be a written opinion of the International Preliminary Examining Authority ("IPEA") except that this does not apply where the applicant chooses an Authority other than this one to be the IPEA and the chosen IPEA has notified the International Bureau under Rule 66.1bis(b) that written opinions of this International Searching Authority will not be so considered.					Authority other		
	If this opinion is, as provided above, considered to be a written opinion of the IPEA, the applicant is invited to submit to the IPEA a written reply together, where appropriate, with amendments, before the expiration of 3 months from the date of mailing of Form PCT/ISA/220 or before the expiration of 22 months from the priority date, whichever expires later.							
	For further options, see Form PCT/IS A/220.							
3.	3. For further details, see notes to Form PCT/ISA/220.							
Name ar	nd maili	ng address of the	ISA/JP	Date of completion of	of this opinion	Authorized	officer	
					-			
Facsimile No.					Telephone :	No.		

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WRITTEN OPINION OF THE INTERNATIONAL SEARCHING AUTHORITY

International application No.

PCT/JP2006/309890

Вох	k No. I	Basis of this opinion	
1.	With	regard to the language, this opinion has been established on the basis of:	
	\boxtimes	the international application in the language in which it was filed	
		the translation of the international application into	which is the language of a
2.		n regard to any nucleotide and/or amino acid sequence disclosed in the international application a ntion, this opinion has been established on the basis of:	nd necessary to the claimed
	a.	type of material	
		a sequence listing	
		table(s) related to the sequence listing	
	b.	format of material	
		on paper	
		in electronic form	
	c.	time of filing/furnishing	
		contained in the international application as filed	
		filed together with the international application in electronic form	
		furnished subsequently to this Authority for the purposes of search	
3.	Ш	In addition, in the case that more than one version or copy of a sequence listing and/or table(s) relative furnished, the required statements that the information in the subsequent or additional copies is identicated or does not go beyond the application as filed, as appropriate, were furnished.	
4.	Addi	itional comments:	

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WRITTEN OPINION OF THE INTERNATIONAL SEARCHING AUTHORITY

International application No.
PCT/JP2006/309890

Box No. V		Reasoned statement under Rule 43bis.1(a)(i) with regard to novelty, inventive step or industrial applicability; citations and explanations supporting such statement			
1.	Statement				
	Novelty (N)	Claims		YES
			Claims	1-13	NO
	Inventive	step (IS)	Claims		YES
			Claims	1-13	NO
	Industrial	applicability (IA)	Claims	1-13	YES
			Claims		NO

2. Citations and explanations:

The following documents have been shown in the ISR.

Document 1: Dinara Daniel et al., Pathway of Apoptosis Induced in Jurkat T Lymphoblasts by Anti-HLA Class I Antibodies, Human Immunology, Vol. 65, pages 189 to 199, 2004 (see, particularly, page 197, left column, line 1 to right column, line 8)

Document 2: Giuliana Cangemi et al., IFN- α mediates the up-regulation of HLA class I on melanoma cells without switching proteasome to immunoproteasome, International Immunology, Vol. 15, No. 12, pages 1415 to 1421, 2003 (see, particularly, page 1416, left column, lines 22 to 24) Document 3: O.R.Burrone et al., Stimulation of HLA-A, B, C by IFN- α . The derivation of Molt 4 variants and the differential expression of HLA-A, B, C subsets, The EMBO Journal, Vol. 4, No. 11, pages 2855 to 2860, 1085 (see, particularly, page 5, Table I.)

Document 4: Naoki Kimura et al., 2D7 diabody bound to the α2 domain of HLA class I efficiently induced caspase-independent cell death against malignant and activated lymphoid cells, Biochemical and Biophysical Research Communications, Vol. 325, pages 1201 to 1209, 2004 Document 5: Laurent Genestier et al., Fas-Independent Apoptosis of Activated T Cells Induced by

Antibodies to the HLA Class I α1 Domain, Blood, Vol. 90, No. 9, pages 3629 to 3639, 1997

Document 1 describes that the anti-HLA class I antibody is useful for inducing cell death to treat the tumor and that since expression of HLA is reduced in the tumor, expression of these tumor tissue compatible conjugates should be restored by a cytokine such as interferon γ .

Accordingly, document 1 can be considered to describe that the interferon is used in combination for enhancing the cell death induction action of the anti-IILA class I antibody when the anti-IILA class I antibody is used as a medicine for inducing cell death to treat the tumor.

Therefore, the subject matters of claims 1-13 of this international application do not appear to be novel or to involve an inventive step in view of document 1.

In this connection, claims 1-13 do not describe that interferon α is used as an interferon and a specific antibody is used as the anti-HLA class I antibody, but documents 2 and 3 each describe that interferon α up-regulates expression of HLA class I on the cell as interferon γ does, and documents 4 and 5 each describe that various kinds of HLA class I antibodies disclosed in the specification of this international application induce cell death in the tumor. Therefore, these aspects are obvious to a person skilled in the art although they are specified in the claims from documents 1-5.

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WRITTEN OPINION OF THE INTERNATIONAL SEARCHING AUTHORITY

International application No.

PCT/JP2006/309890

Supplemental Box					
In case the space in any of the preceding boxes is not sufficient. Continuation of: Int.Cl.					
A61P17/06(2006.01),	A61P21/04(2006.01),				
A61P25/00(2006.01),	A61P29/00(2006.01),				
A61P35/00(2006.01),	A61P35/02(2006.01),				
A61P37/02(2006.01),	A61P43/00(2006.01)				